## INTERNATIONAL SEARCH REPORT

International Application No PC1 03/09991

A. CLASSIFICATION OF SUBJECT MATTER IPC 7 G01N33/53 C12Q1/68

According to International Patent Classification (IPC) or to both national classification and IPC

## B. FIELDS SEARCHED

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

## EPO-Internal

5

2

Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
Χ .	BRIAN STAUSBOL-GRON ET AL: "De novo identification of cell-type specific antibody-antigen pairs by phage display subtraction. Isolation of a human single chain antibody fragment against human keratin 14" EUR. J. BIOCHEM., vol. 268, 2001, pages 3099-3107, XP002267098 the whole document	1-30
X	WO 00 52054 A (GENENTECH INC) 8 September 2000 (2000-09-08) page 3, line 6 - line 11 page 18, line 5 - line 35 page 9, line 20 - line 30 claims 5,6	1-30

X Further documents are listed in the continuation of box C.	Patent family members are listed in annex.				
"A" document defining the general state of the art which is not considered to be of particular relevance  "E" earlier document but published on or after the international filing date  "L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)  "O" document referring to an oral disclosure, use, exhibition or other means  "P" document published prior to the international filing date but later than the priority date claimed	"T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention  "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone  "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art.  "&" document member of the same patent family				
Date of the actual completion of the international search  16 January 2004	Date of mailing of the international search report  1 0. 02 2004				
Name and mailing address of the ISA  European Patent Office, P.B. 5818 Patentlaan 2  NL - 2280 HV Rijswijk  Tel. (+31-70) 340-2040, Tx. 31 651 epo nl,  Fax: (+31-70) 340-3016	Authorized officer  MALIN SÖDERMAN /EÖ				

Form PCT/ISA/210 (second sheet) (July 1992)

# INTERNATIONAL SEARCH REPORT

International Application No PC 03/09991

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	lation) DOCUMENTS CONSIDERED TO BE RELEVANT		
Category °	Citation of document, with indication, where appropriate, of the relevant passages		Relevant to daim No.
X	BRIAN STAUSBOL-GRON ET AL: "A model phage display subtraction method with potential for analysis of differential gene expression" FEBS LETTERS, vol. 391, 1996, pages 71-75, XP002267099 the whole document		1-30
X	WO 02 39120 A (BURIONI ROBERTO ;GEORGE SHAJI T (US); BIONOVA PHARMACEUTIALS INC () 16 May 2002 (2002-05-16) page 5, line 54 -page 8, line 55 page 16, line 3 - line 32 page 21, line 52 -page 23, line 30 page 26, line 38 - line 47		1-30
A	WLODEK MANDECKI ET AL: "A Mathematical Model for Biopanning (Affinity Selection) Using Peptide Libraries on Filamentous Phage"  J. THEOR. BIOL., vol. 176, 1995, pages 523-530, XP002267100 the whole document		1-30
A	BRENT R. WILLIAMS ET AL: "Polyclonal anti-colorectal cancer Fab phage display library selected in one round using density gradient centrifugation to separate antigenbound and free phage" IMMUNOLOGY LETTERS, vol. 81, 2002, pages 141-148, XP002267101 the whole document		1-30
			at.

# INTERNATIONAL SEARCH REPORT

marmation on patent family members

PC 03/09991

Patent document cited in search report		Publication Patent family date member(s)		Publication date	
WO 0052054	A	08-09-2000	AU CA EP JP WO US	3389800 A 2361877 A1 1157041 A2 2002543044 T 0052054 A2 2003219434 A1	21-09-2000 08-09-2000 28-11-2001 17-12-2002 08-09-2000 27-11-2003
WO 0239120	Α	16-05-2002	AU WO	2023602 A 0239120 A1	21-05-2002 16-05-2002

These Notes are intended to give the basic instructions concerning the filing of amendments under article 19. The Notes are based on the requirements of the Patent Cooperation Treaty, the Regulations and the Administrative Instructions under that Treaty. In case of discrepancy between these Notes and those requirements, the latter are applicable. For more detailed information, see also the PCT Applicant's Guide, a publication of WIPO.

In these Notes, "Article", "Rule", and "Section" refer to the provisions of the PCT, the PCT Regulations and the PCT Administrative Instructions, respectively.

#### INSTRUCTIONS CONCERNING AMENDMENTS UNDER ARTICLE 19

The applicant has, after having received the international search report, one opportunity to amend the claims of the international application. It should however be emphasized that, since all parts of the international application (claims, description and drawings) may be amended during the international preliminary examination procedure, there is usually no need to file amendments of the claims under Article 19 except where, e.g. the applicant wants the latter to be published for the purposes of provisional protection or has another reason for amending the claims before international publication. Furthermore, it should be emphasized that provisional protection is available in some States only.

#### What parts of the international application may be amended?

Under Article 19, only the claims may be amended.

During the international phase, the claims may also be amended (or further amended) under Article 34 before the International Preliminary Examining Authority. The description and drawings may only be amended under Article 34 before the International Examining Authority.

Upon entry into the national phase, all parts of the international application may be amended under Article 28 or, where applicable, Article 41.

#### When?

Within 2 months from the date of transmittal of the international search report or 16 months from the priority date, whichever time limit expires later. It should be noted, however, that the amendments will be considered as having been received on time if they are received by the International Bureau after the expiration of the applicable time limit but before the completion of the technical preparations for international publication (Rule 46.1).

#### Where not to file the amendments?

The amendments may only be filed with the International Bureau and not with the receiving Office or the International Searching Authority (Rule 46.2).

Where a demand for international preliminary examination has been/is filed, see below.

#### How?

Either by cancelling one or more entire claims, by adding one or more new claims or by amending the text of one or more of the claims as filed.

A replacement sheet must be submitted for each sheet of the claims which, on account of an amendment or amendments, differs from the sheet originally filed.

All the claims appearing on a replacement sheet must be numbered in Arabic numerals. Where a claim is cancelled, no renumbering of the other claims is required. In all cases where claims are renumbered, they must be renumbered consecutively (Administrative Instructions, Section 205(b)).

The amendments must be made in the language in which the international application is to be published.

## What documents must/may accompany the amendments?

#### Letter (Section 205(b)):

The amendments must be submitted with a letter.

The letter will not be published with the international application and the amended claims. It should not be confused with the "Statement under Article 19(1)" (see below, under "Statement under Article 19(1)").

The letter must be in English or French, at the choice of the applicant. However, if the language of the international application is English, the letter must be in English; if the language of the international application is French, the letter must be in French.

The letter must indicate the differences between the claims as filed and the claims as amended. It must, in particular, indicate, in connection with each claim appearing in the international application (it being understood that identical indications concerning several claims may be grouped), whether

- the claim is unchanged;
- (ii) the claim is cancelled;
- (iii) the claim is new;
- (iv) the claim replaces one or more claims as filed;
- (v) the claim is the result of the division of a claim as filed.

# The following examples illustrate the manner in which amendments must be explained in the accompanying letter:

- [Where originally there were 48 claims and after amendment of some claims there are 51]:
   "Claims 1 to 29, 31, 32, 34, 35, 37 to 48 replaced by amended claims bearing the same numbers;
   claims 30, 33 and 36 unchanged; new claims 49 to 51 added."
- [Where originally there were 15 claims and after amendment of all claims there are 11]:
   "Claims 1 to 15 replaced by amended claims 1 to 11."
- 3. [Where originally there were 14 claims and the amendments consist in cancelling some claims and in adding new claims]:
  - "Claims 1 to 6 and 14 unchanged; claims 7 to 13 cancelled; new claims 15, 16 and 17 added." or "Claims 7 to 13 cancelled; new claims 15, 16 and 17 added; all other claims unchanged."
- 4. [Where various kinds of amendments are made]: "Claims 1-10 unchanged; claims 11 to 13, 18 and 19 cancelled; claims 14, 15 and 16 replaced by amended claim 14; claim 17 subdivided into amended claims 15, 16 and 17; new claims 20 and 21 added."

#### "Statement under article 19(1)" (Rule 46.4)

The amendments may be accompanied by a statement explaining the amendments and indicating any impact that such amendments might have on the description and the drawings (which cannot be amended under Article 19(1)).

The statement will be published with the international application and the amended claims.

# It must be in the language in which the international application is to be published.

It must be brief, not exceeding 500 words if in English or if translated into English.

It should not be confused with and does not replace the letter indicating the differences between the claims as filed and as amended. It must be filed on a separate sheet and must be identified as such by a heading, preferably by using the words "Statement under Article 19(1)."

It may not contain any disparaging comments on the international search report or the relevance of citations contained in that report. Reference to citations, relevant to a given claim, contained in the international search report may be made only in connection with an amendment of that claim.

# Consequence if a demand for international preliminary examination has already been filed

If, at the time of filing any amendments and any accompanying statement, under Article 19, a demand for international preliminary examination has already been submitted, the applicant must preferably, at the time of filing the amendments (and any statement) with the International Bureau, also file with the International Preliminary Examining Authority a copy of such amendments (and of any statement) and, where required, a translation of such amendments for the procedure before that Authority (see Rules 55.3(a) and 62.2, first sentence). For further information, see the Notes to the demand form (PCT/IPEA/401).

## Consequence with regard to translation of the international application for entry into the national phase

The applicant's attention is drawn to the fact that, upon entry into the national phase, a translation of the claims as amended under Article 19 may have to be furnished to the designated/elected Offices, instead of, or in addition to, the translation of the claims as filed.

For further details on the requirements of each designated/elected Office, see Volume II of the PCT Applicant's Guide.

# **PATENT COOPERATION TREATY**



# INTERNATIONAL SEARCH REPORT

(PCT Article 18 and Rules 43 and 44)

Applicant's or agent's file reference FOR FURTHER see Notification of Transmittal of International Search Report (Form PCT/ISA/220) as well as, where applicable, item 5 below.						
BIOT/ P29002PC	ACTION "					
International application No.	International filing date (day/month/year)		(Earliest) Priority Date (day/month/year)			
PCT/EP 03/09991	03/09/2003		04/09/2002			
Applicant				·		
BIOINVENT INTERNATIONAL AF	3 					
This International Search Report has beer according to Article 18. A copy is being tra	n prepared by this International Insmitted to the International Bu	Searching Author ureau.	ity and is transmitt	ed to the applicant		
g acceptant g acceptant g						
This International Search Report consists						
X It is also accompanied by	a copy of each prior art docum	ent cited in this re	port.			
Basis of the report				<u> </u>		
a With regard to the language, the i	nternational search was carried	d out on the basis	of the international	application in the		
language in which it was filed, unli	ess otherwise indicated under t	his item.				
the international search w Authority (Rule 23.1(b)).	as carried out on the basis of a	translation of the	international applic	cation furnished to this		
b. With regard to any nucleotide an	d/or amino acid sequence dis	closed in the inter	national application	n, the international search		
was carried out on the basis of the	sequence listing :					
	nal application in written form. rnational application in compute	er readable form				
1	this Authority in written form.	or roadable form.				
	this Authority in computer read	lble form.				
the statement that the sub	the statement that the subsequently furnished written sequence listing does not go beyond the disclosure in the international application as filed has been furnished.					
	rmation recorded in computer r	readable form is id	lentical to the writte	en sequence listing has been		
2. Certain claims were four	nd unsearchable (See Box I).					
3. Unity of invention is lack	•					
4. With regard to the title,				•		
the text is approved as su						
ا اسما	ned by this Authority to read as		777777777			
METHOD FOR SCREENING A	NTI-LIGAND LIBRARI	ES FOR IDE	NITEXING AND	TI-LIGANDS SPECIFIC		
FOR DIFFERENTIALLY AND	INFREQUENTLY EXPR	ESSED LIGAN	NDS			
5. With regard to the abstract,						
X the text is approved as sul				w <del></del>		
the text has been establish within one month from the	ned, according to Rule 38.2(b), date of mailing of this internation	by this Authority a onal search report	is it appears in Box , submit comments	to this Authority.		
6. The figure of the <b>drawings</b> to be publi			5			
X as suggested by the applic				None of the figures.		
because the applicant faile	ed to suggest a figure.		<del></del>			
because this figure better	characterizes the invention.					



Box I	Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)
This Inte	ernational Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:
1. χ	Claims Nos.: 28 because they relate to subject matter not required to be searched by this Authority, namely: see FURTHER INFORMATION sheet PCT/ISA/210
2. X	Claims Nos.:  1, 2, 24, 25 because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:  See FURTHER INFORMATION sheet PCT/ISA/210
3.	Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).
Box il	Observations where unity of invention is lacking (Continuation of Item 2 of first sheet)
This Inte	ernational Searching Authority found multiple inventions in this international application, as follows:
1.	As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
2	As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3.	As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:
4.	No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:
Remark	The additional search fees were accompanied by the applicant's protest.  No protest accompanied the payment of additional search fees.

## FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

Continuation of Box I.1

Claims Nos.: 28

Claim 28 relate to methods of treatment of the human or animal body by surgery or by therapy or diagnostic methods practised on the human or animal body (PCT Rule 39.1(iv)). Nevertheless, a search has been executed for this claim. The search has been based on the alleged effects of the compounds or compositions.

Continuation of Box I.2

Claims Nos.: 1, 2, 24, 25

The claim or claims shall define the matter for which protection is sought. Claims shall be clear and concise, see Article 6 PCT. Claim 25 contains references to the description and the drawings. According to Rule 6.2(a) PCT, claims should not contain such references except where absolutely necessary, which is not the case here.

The claim or claims shall define the matter for which protection is sought. Claims shall be clear and concise, see Article 6 PCT. In claim 1 and 2 step (iv) the phrases "A, B, C & D = are the participants..." and "a, b, c & d = are the coefficients..." are not clear and should be amended.

Claim 24 does not meet the requirements of Article 6 PCT in that the matter for which protection is sought is not clearly defined. The claim attempts to define the subject-matter in terms of the result to be achieved which merely amounts to a statement of the underlying problem. The technical features necessary for achieving this result should be added. It is not clear which stimulus is added and what effect is has on the expression of the target ligands.

The applicant's attention is drawn to the fact that claims, or parts of claims, relating to inventions in respect of which no international search report has been established need not be the subject of an international preliminary examination (Rule 66.1(e) PCT). The applicant is advised that the EPO policy when acting as an International Preliminary Examining Authority is normally not to carry out a preliminary examination on matter which has not been searched. This is the case irrespective of whether or not the claims are amended following receipt of the search report or during any Chapter II procedure.